The Impact of Probiotic Supplementation on Weight and Body Composition in Older Adults: A Systematic Review

Zahra Hoseini-Tavassol¹, Hanieh-Sadat Ejtahed^{1,2,*}, Mohammad Javad Mansourzadeh³, Fateme Ettehad-Marvasti¹, Golnaz Bahramali⁴, Kazem Khalagi^{1,3}, Shirin Hasani-Ranjbar^{1,*}, Seyed-Davar Siadat⁵, Iraj Nabipour⁶, Afshin Ostovar^{3,7}, Bagher Larijani²

Submitted: 29 April 2024 Revised: 24 May 2024 Accepted: 29 May 2024 Published: 1 July 2024

Background: Obesity as a complex disease significantly impacts life quality and expectancy in older adults. Managing obesity in older adults is particularly challenging. The use of probiotics has been recommended as a potential strategy for weight management. So, this study aims to review the effectiveness of probiotics on weight and body composition in older adults. Methods: A systematic search was conducted across multiple databases including PubMed, Web of Science, Scopus, and Embase, to retrieve relevant publications up to May 2023. Eligibility criteria were applied to select clinical trials that reported the effects of

to retrieve relevant publications up to May 2023. Eligibility criteria were applied to select clinical trials that reported the effects of probiotics on weight and body composition in older adults. Two independent investigators performed data extraction according to a pre-designed table and assessed the quality of the selected studies using the Cochrane risk-of-bias tool.

Results: A total of 705 records were found, and 12 articles were considered for the review. In four studies, the use of probiotics has been demonstrated to decrease on weight and body composition in older adults. However, due to the high diversity of bacterial species and multispecies probiotics used, it cannot be concluded which type of probiotics is recommended for managing obesity in older adults.

Conclusions: While there is evidence suggesting that probiotics may have an impact on weight management, the effectiveness of probiotics on obesity in older adults is a more complex issue and influenced by various factors like aging, underlying diseases and lifestyle. More clinical studies are needed to determine the effective combination of probiotics for managing obesity in older adults.

Keywords: obesity; probiotic; older adults; body weight; body composition

Introduction

Obesity is a complex disease that is associated with a range of comorbidities, including cardiovascular disease, gastrointestinal disorders, type 2 diabetes, musculoskeletal disorders, respiratory difficulties, and psychological problems. Obesity can have a significant impact on life quality and expectancy and, increase the risk of mortality [1,2]. Today, there is growing concern about the potential burden of obesity prevalence in the older adult population worldwide [3]. Remarkably obesity management strategies for older adults are more challenging for various reasons [4].

Therefore, it is important to address obesity in older adults and develop appropriate clinical approaches to manage this problem.

Past study has shown the impact of gut microbiota on obesity among older adults. The composition of the gut microbiota is associated with obesity in these individuals [5]. Available evidence indicates that the use of probiotics is one of the recommended strategies for weight management [6]. Although depending on individual status and requirements, probiotics use should be determined according to the age and health conditions of the person [7].

¹Obesity and Eating Habits Research Center, Endocrinology and Metabolism Clinical Sciences Institute, Tehran University of Medical Sciences, 1417653761 Tehran, Iran

²Endocrinology and Metabolism Research Center, Endocrinology and Metabolism Clinical Sciences Institute, Tehran University of Medical Sciences, 1417653761 Tehran, Iran

³Osteoporosis Research Center, Endocrinology and Metabolism Clinical Sciences Institute, Tehran University of Medical Sciences, 1417653761 Tehran, Iran

⁴Department of Hepatitis and AIDS and Blood Borne Diseases, Pasteur Institute of Iran, 1316943551 Tehran, Iran

⁵Microbiology Research Center, Pasteur Institute of Iran, 1316943551 Tehran, Iran

⁶The Persian Gulf Marine Biotechnology Research Center, The Persian Gulf Biomedical Sciences Research Institute, Bushehr University of Medical Sciences, 7514633341 Bushehr, Iran

⁷Department of Epidemiology and Biostatistics, School of Public Health, Tehran University of Medical Sciences, 1417653761 Tehran, Iran

^{*}Correspondence: haniejtahed@yahoo.com; ejtahed-h@sina.tums.ac.ir (Hanieh-Sadat Ejtahed); shirinhasanir@yahoo.com; sh_hasani@tums.ac.ir (Shirin Hasani-Ranjbar)

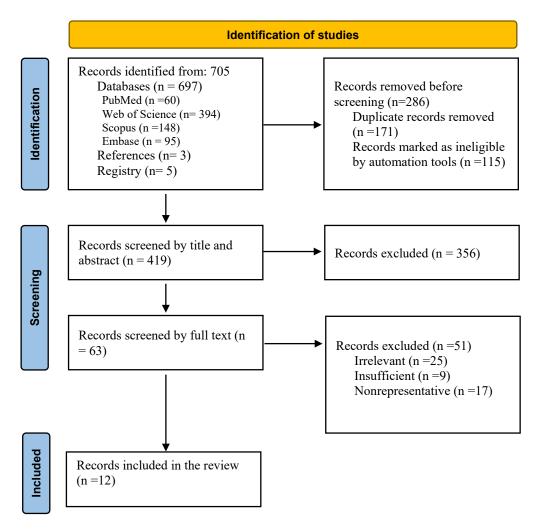


Fig. 1. Flow diagram of the study selection process.

Exploring the potential of probiotics as biotherapeutic candidates for managing obesity in older adults shows promise for future research. So, our systematic review aims to evaluate the effectiveness of probiotic interventions compared to placebo or other interventions on weight and body composition in older adults.

Methods

Study Design and Search Strategy

This systematic review followed the Preferred Reporting Items for Systematic reviews and Meta-Analyses (PRISMA) guidelines for conducting and reporting the study results [8] (Supplementary Table 1). A systematic search was conducted using various databases such as PubMed, Web of Science, Scopus, and Embase to retrieve publications until May 2023. The search focused on the concepts of Microbiota, Obesity, and Older adults. The search strategy and terms used are shown in Supplementary Table 2. The results from all databases were imported and collected in Zotero reference management software version 6.0.27 (Digital Scholar, Vienna, VA, USA),

and duplicates were removed using software and manual inspection by two independent investigators. Additionally, the reference lists of the included studies were screened, and the International Clinical Trials Registry Platform (ICTRP) (https://trialsearch.who.int) was searched to find more eligible records.

Eligibility Criteria and Study Selection

The retrieved publications were screened based on titles and abstracts, and two investigators excluded irrelevant studies. The inclusion criteria were clinical trials that reported the effects of probiotics on weight and body composition in older adults with any health condition and studies with full-text articles available in English. The exclusion criteria were animal, observational and ecological studies, review articles, editorials, letters, comments and studies with insufficient data.

	Author-date	Country	Sample size	Intervention group characteristics	Control group characteristics	Women percentage	Participants' underlying condition	Probiotic supplement and amount	Co-supplementation	Control group Supplement	Intervention duration	Anthropometri measurements change
1	João Valentini Neto et al. (2013) [10]	Brazil	17	n = 9 67.9 ± 4.5 y	$n = 8$ $67.9 \pm 4.5 \text{ y}$	76.5%	Overweight	L. paracasei, L. rhamnosus, L. acidophilus, B. lactis 10 ⁸ to 10 ⁹	Fructooligosaccharide 6 g	Maltodextrin placebo	3 months	NS
2	Fabíola Málaga Barreto <i>et al.</i> (2014) [11]	Brazil	24	n = 12, 62 y	n = 12, 63 y	100%	Overweight, Metabolic syndrome, Postmenopause	L. plantarum (Lp 115) 1.25 × 10 ⁷ UFC/g, Daily: 80 mL	Fermented milk with Probiotic	Non-fermented milk	90 days	NS
3	Kerry L Ivey et al. (2014) [12]	Australia	156	n (Probiotic yoghurt) = 40	n (Probiotic yoghurt) = 37	100%	Obesity	L. acidophilus La5,	Probiotic yoghurt	Probiotic yoghurt + Placebo	6 weeks	NS
				$68 \pm 8 \text{ y}$ $n \text{ (Control milk)} = 39$ $65 \pm 7 \text{ y}$	$68 \pm 8.7 \text{ y}$ n (Control milk) = 40 $65 \pm 8 \text{ y}$			B. animalis subsp. lactis Bb12 3.0×10^9 CFU/d	OR Control milk	OR Control milk + Placebo		
4	Lena K. Brahe <i>et al.</i> (2015) [13]	Denmark	58	$n = 19$ 61.4 ± 6.5	$n = 19$ 60.6 ± 6.4 And $n = 16$ 58.5 ± 5.3	100%	Obesity, Postmenopause	<i>L. paracasei F19</i> 9.4 × 10 ¹⁰		Flaxseed mucilage 10 g And Placebo	6 weeks	NS
	Iaryam Tajabadi- Ebrahimi <i>et al</i> . (2017) [14]	Iran	60	$n = 30,$ $64.2 \pm 12.0 \text{ y}$	$n = 30, 64.0 \pm 11.7$	у	Overweight, Type 2 Diabetes, Coronary Heart Disease	L. acidophilus 2×10^9 CFU/g, L. casei 2×10^9 CFU/g, B. bifidum 2×10^9 CFU/g	Inulin 800 mg	Starch placebo	12 weeks	NS
6	Fariba Raygan <i>et al</i> . (2018) [15]	Iran	60	n = 30	n = 30	50%	Overweight, Diabetes, Coronary heart disease	$L.\ acidophilus,$ $B.\ bifidum,$ $L.\ reuteri,$ $L.\ fermentum\ each\ 2\times10^9$	Vitamin D3 50,000 IU every 2 weeks	Placebo	12 weeks	NS
7	Monika Szulińska <i>et al.</i> (2018) [16]	Poland	71	$n = 24$, $58.72 \pm 7.25 \text{ y}$	n (Low-dose probiotic) = 24 56.38 ± 6.55 y, n (High-dose probiotic) = 23 55.16 ± 6.87 y	100%	Obesity, Postmenopause	B. bifidum W23, B. lactis W51, B. lactis W52, L. acidophilus W37, L. brevis W63, L. casei W56, L. salivarius W24, L. lactis W19, L. lactis W58 Low dose = 2.5 × 10 ⁹ CFU High dose = 1 × 10 ¹⁰ CFU		Placebo	12 weeks	↓ WC ↓ Fat

Table 1. Continued.											
Author-date	Country	Sample size	Intervention group characteristics	Control group characteristics		Participants' underlying condition	Probiotic supplement and amount	Co-supplementation	Control group Supplement	Intervention duration	Anthropometric measurements change
Raffaella 8 Cancello <i>et al</i> . (2019) [17]	Italy	20	$n = 20,$ $79.1 \pm 3.3 \text{ y}$	$n = 20,$ $79.1 \pm 3.3 \text{ y}$	100%	Obesity, Postmenopause Somewhat: Arthrosis, Hypertension, Obstructive sleep apnea, Type 2 diabetes	Treptococcus Thermophilus, B. breve,	Mediterranean diet administered	Mediterranean diet administered	2 weeks	↓ Weight
						1 7 31	B. longum, B. infantis, L. acidophilus, L. plantarum, L. paracasei, L. delbrueckii bulgaricus				
Alireza 9 Farrokhian <i>et a</i> (2019) [18]	l. Iran	60	n = 30 $64.2 \pm 12.0 \text{ y}$	n = 30 64.0 ± 11.7	64%	Overweight, Diabetes, Coronary Heart Disease	L. acidophilus strain T16, L. casei strain T2, B. bifidum strain T1 2 × 10 ⁹ CFU/g	Inulin 800 mg	Placebo	12 weeks	NS
Mahsa Raji 10 Lahiji <i>et al.</i> (2021) [19]	Iran	72	n = 36 56.5 ± 5.0	n = 36 58.3 ± 6.4	100%	Overweight, Obesity, Postmenopause, Breast cancer survivor	$\begin{array}{c} \textit{L. casei } 1.5 \times 10^9 \text{ CFU/g,} \\ \textit{L. acidophilus } 1.5 \times 10^{10} \text{ CFU/g,} \\ \textit{L. rhamnosus } 3.5 \times 10^9 \text{ CFU/g,} \\ \textit{L. sbulgaricus } 2.5 \times 10^8 \text{ CFU/g,} \\ \textit{B. breve } 1 \times 10^{10} \text{ CFU/g,} \\ \textit{B. longum } 5 \times 10^8 \text{ CFU/g,} \\ \textit{Streptococcus Thermophilus} \\ 1.5 \times 10^8 \text{ CFU/g} \end{array}$	Fructooligosaccharide 35 mg	Placebo	8 weeks	NS
Chaiyavat 11 Chaiyasut <i>et al</i> (2022) [20]	. Thailand	48	$n = 24$ $61.63 \pm 0.84 \text{ y}$	$n = 24$ $58.79 \pm 1.21 \text{ y}$	79%		L. paracasei HII01 2×10^{10} CFU, B. breve 2×10^{10} CFU, B. longum 1×10^{10} CFU		Corn starch placebo	12 weeks	↓ Weight ↓ BMI ↓ WC ↓ HC
Mariusz 12 Kaczmarczyk et al. (2022) [2		56	Group 1: 18 54 ± 7 y Group 2: 18 56 ± 7 y	20 58 ± 8 y	100%	Obesity, Postmenopause	B. bifidum W23, B. lactis W51, B. lactis W52, L. acidophilus W37, L. brevis W63, L. casei W56, Ligilactobacillus salivarius W24, L. lactis W19, L. lactis W58 Group 1: 2.5 × 10 ⁹ CFU/day Group 2: 1 × 10 ¹⁰ CFU/day		Placebo	12 weeks	↓ Weight ↓ BMI

y, year.

L., Lactobacillus; B., Bifidobacterium.

BMI, body mass index; WC, waist circumference; HC, hip circumference.

NS: No significant differences in terms of anthropometric measurements.



Data Extraction

The full-text articles of potentially relevant studies that met the inclusion criteria were reviewed. The extracted information included the Author-Date, Country, Sample size, Intervention group characteristics, Control group characteristics, Women percentage, Participants' underlying condition, Probiotic Supplement and amount, Cosupplementation, Control group Supplement, Intervention duration and Anthropometric measurements change, which summarized in a pre-designed data extraction table by two independent investigators. Any disagreements were resolved by consulting with the principal investigator.

Quality Assessment

The Cochrane risk-of-bias tool for randomized trials was used for the quality assessment of included studies [9]. It assesses bias in several domains, and the answers to signalling questions lead to judgments of "Low risk of bias", "Unclear risk of bias" or "High risk of bias". The blinded quality assessment of each study was performed by two investigators. Any dissimilarities were settled through discussion or consensus between the two investigators.

Result

A total of 705 records were found through searches. After removing 171 duplicate records and 115 records marked as ineligible by automation tools, the titles and abstracts of 419 articles were screened. Out of those, 356 articles were excluded based on predefined exclusion criteria, or irrelevant to the main topic. After screening the 63 full-texts, ineligible articles, studies with no relevant or insufficient outcomes were excluded, and finally, 12 studies were considered for the review [10–21] (Fig. 1). Also, a clinical trial with a 'Not recruiting' status was found on the ICTRP that may be eligible for inclusion [22].

The results of the included articles' risk-of-bias assessment are presented in Fig. 2 (detailed in **Supplementary Table 3**).

This review showed that a limited number of clinical studies have reported the effect of probiotics on obesity in older adults. Table 1 (Ref. [10–21]) shows the data of the articles that reported the effect of various probiotics on weight and body composition in older adults or late middle-aged populations.

Four studies were conducted in Iran, two in Brazil, and the others in Australia, Denmark, Poland, Italy, Spain and Thailand. The sample sizes of the studies varied from 17 to 156 participants, and total sample size was 702. The mean age of the participants in these studies has been between 54 and 79.1 years and most of them are women. Participants' underlying condition includes somewhat: Obesity, Overweight, Metabolic syndrome, Postmenopause, Diabetes, Coronary Heart Disease, Arthrosis, Hypertension, Obstructive sleep apnea and Breast cancer survivor. All studies,

except one, have used probiotic mixtures as an intervention. The Lactobacillus (11 studies) [10–18,20,21] and Bifidobacterium (8 studies) [12,14–18,20,21] were the most abundant genera used as probiotics in these studies. One study has used Streptococcus Thermophilus [17], and one study has used Ligitactobacillus salivarius [21]. The multispecies probiotic supplement has been used in 10 studies [10,12,14-21]. The probiotic amount varied in different studies, with a minimum dose of 10⁸ and a maximum dose of 9.4×10^{10} . Four studies used synbiotics, Fructooligosaccharide [10,19] and Inulin [14,18] as an intervention in their randomized clinical trials. In one study, the Mediterranean diet was administered alongside probiotic consumption [17]. Two studies use fermented milk with probiotics [11] and probiotic yogurt/control milk and probiotics [12] as interventions. In one study, vitamin D was also prescribed alongside probiotic consumption [15]. The intervention duration was between 2 weeks and 3 months. Two studies demonstrated the alteration of gut microbiota composition before and after intervention [13,17].

In four studies, the probiotic intervention has been shown to have a body composition-reducing effect and there was a reduction in anthropometric indices, weight, body mass index (BMI), waist circumference (WC), hip circumference (HC), and fat in the studies [16,17,20,21]. These studies use *Bifidobacterium bifidum W23*, *B. lactis W51*, *B. lactis W52*, *Lactobacillus acidophilus W37*, *L. brevis W63*, *L. casei W56*, *L. salivarius W24*, *L. lactis W19* and *L. lactis W58* [16,21], *B. breve*, *B. longum*, *L. paracasei* [17,20], *Treptococcus Thermophilus*, *B. infantis*, *L. acidophilus*, *L. plantarum*, *L. delbrueckii bulgaricus* [17] as multispecies probiotic supplements.

Discussion

Few studies are reporting the effectiveness of probiotics on weight, anthropometric indices, and other factors related to obesity. Therefore, the results of this review study do not provide accurate information about probiotics' effectiveness in older adults. Despite this, numerous studies have demonstrated the impact of probiotics in managing obesity [23–27], various factors influence the effect of probiotics on obesity in older adults, such as aging, underlying diseases [28], dietary habits [29], and physical activity [30]. Furthermore, the use of multispecies probiotic supplements complicates the identification of effective probiotic types and mechanisms in the included studies. Therefore, in some studies, the use of probiotics has a significant effect on weight and body composition, while in others, it does not. Also, since most of the study participants are post-menopausal women, the effectiveness of probiotics may be partially related to this process [31]. So, this subject is highly challenging to study and it cannot be said which type of probiotics is recommended for managing obesity in older adults.

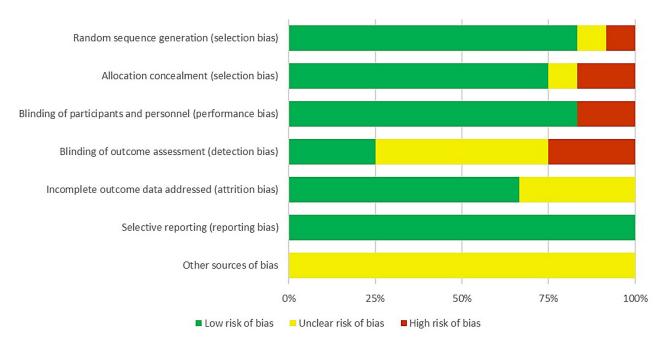


Fig. 2. The results of the included articles' risk-of-bias assessment.

The results of this study showed that *Lactobacillus* and *Bifidobacterium* are the most common genera used as probiotics. In previous studies, the effect of these genera as a probiotic on obesity has been investigated [31].

In Crovesy et al.'s [32] study Lactobacillus plantarum and L. rhamnosus, L. curvatus, L. gasseri, L. amylovorus, L. acidophilus and L. casei, and multiple species supplement of Lactobacillus effect is highlighted. In Ejtahed et al.'s [33] systematic review, they showed that some specific strains of Lactobacillus and Bifidobacterium species including Lactobacillus casei, L. rhamnosus, L. gasseri, Lactobacillus plantarum, B. infantis, B. longum and B. breve showed an anti-obesity effect or decreased in weight, waist circumference or fat mass in most animal and human studies. In previous clinical trial studies and systematic reviews Lactobacillus gasseri [34], L. rhamnosus [35], L. plantarum [36,37], L. curvatus [37] with or without other species of Lactobacillus and Bifidobacterium association have a potential effect in weight and body composition reduction [25], but we could not confirm the effectiveness of these species in the included studies. Three articles in which weight loss has been observed, used Lactobacillus acidophilus in their multispecies probiotics supplement, but there is disagreement about the effectiveness of this species in obesity [25]. This functional difference at the species level of Lactobacillus is likely due to genetic variation involved in lipid and carbohydrate metabolism [38].

Regarding the effects of *Bifidobacterium* as a probiotic on obesity, commonly used strains in previous studies are *B. animalis*, *B. bifidum*, *B. lactis* and *B. breve* [25]. These studies show positive effects of *Bifidobacterium lactis* Bb-12 [39] and *B. animalis* ssp. *Lactis* [40–42] *B. bifidum*, *B. longum* [43], *B. breve* [44] on anthropometric in-

dices reduction. However, future clinical trials are necessary to further investigate these effects.

Generally, *Lactobacillus* and *Bifidobacterium* potential mechanisms of action include maintenance intestinal barrier, reduction of inflammation, production of SCFAs like linoleic, propionic, acetic, and butyric acids, improving glucose and lipid metabolism, modulation of energy homeostasis and regulation of appetite [45].

Furthermore, probiotics have the potential to modify the composition of the gut microbiota and alleviate dysbiosis, which could aid in the management of obesity. This indicates that the utilization of probiotics might play a role in controlling weight by influencing the gut microbiota [31]. Here, changes in gut microbiota after probiotic consumption are seen in two studies [13,17]. The Crovesy *et al.*'s [46] systematic review found a higher relative abundance of the genus *Lactobacillus* and *Bifidobacterium* in obese individuals compared to non-obese individuals. Such data affect the role of probiotics in obesity by changing the composition of microbiota and it seems that these changes as well as the effective mechanisms should be investigated at the species level.

In Michael et al.'s [47] study, 6-month supplementation with a probiotic supplement contained Lactobacillus acidophilus, L. acidophilus, L. plantarum, Bifidobacterium bifidum and B. animalis subsp. lactis resulted in significant weight reduction and improved well-being. Also in another study, they supplemented overweight adults with the same probiotic supplement for 9 months and found a significant reduction in body weight without any lifestyle restrictions [48]. This highlights the effect of extended intervention duration that wasn't observed in our included studies.



While conducting this systematic review, we encountered some limitations. Most of the studies included in this review were not aimed at assessing the effect of probiotics on obesity or anthropometric indices in older adults. Also, the underlying conditions of the participants in these studies have been diverse, which may impact the efficacy of probiotics. Additionally, important factors such as nutritional status [49] and co-administration of probiotics with other medications [50] may not have been adequately addressed in these studies. In this study, articles were reviewed, considering the different definitions that each study had considered for old age, as well as the background conditions of individuals such as menopause in women. So, the study results have a lot of heterogeneity. Therefore, due to the lack of sufficient evidence, it is unlikely that our question will be answered. Considering that probiotics are also being studied to see if they can help older adults stay healthy, improve their quality of life and slow down the aging process [51-53], more clinical studies are needed to determine the effective combination and dosage of probiotics for managing obesity in older adults, considering their underlying conditions like chronic diseases, medications, and lifestyle.

Conclusions

In conclusion, the effectiveness of probiotics in managing weight and obesity in older adults is a complex and challenging topic to study, due to the diverse underlying conditions and factors influencing their efficacy. However, several studies have demonstrated the impact of probiotics in managing obesity, with *Lactobacillus* and *Bifidobacterium* being the most common genera used as probiotics. While there is evidence suggesting that probiotics may have an impact on weight management, more research is needed to fully understand their effectiveness and mechanisms in older adults.

Availability of Data and Materials

All data generated in this study are supported by the published articles in PubMed/Medline, Web of Science, Scopus and Embase databases, and all data generated are included in this published article.

Author Contributions

ZHT, HSE: Investigation, conceptualization, designing, writing — original draft. MJM and FEM: Investigation, designing, writing — review & editing. GB and KK: Investigation, writing — review & editing. SHR and HSE: Conceptualization, designing, project administration, supervision, writing — review & editing. SHR, SDS, IN, AO and BL: Conceptualization, writing — review & editing. All authors reviewed and approved the final manuscript. All authors have participated sufficiently in the work and are accountable for all aspects of the study.

Ethics Approval and Consent to Participate

Not applicable.

Acknowledgment

Not applicable.

Funding

This study was funded by the Endocrinology and Metabolism Research Institute, Tehran University of Medical Sciences, Tehran, Iran (Number: 1401-3-91-1213).

Conflict of Interest

The authors declare no conflict of interest.

Supplementary Material

Supplementary material associated with this article can be found, in the online version, at https://doi.org/10.23812/j.biol.regul.homeost.agents.20243807.431.

References

- [1] Fruh SM. Obesity: Risk factors, complications, and strategies for sustainable long-term weight management. Journal of the American Association of Nurse Practitioners. 2017; 29: S3–S14.
- [2] Lin X, Li H. Obesity: Epidemiology, Pathophysiology, and Therapeutics. Frontiers in Endocrinology. 2021; 12: 706978.
- [3] Decaria JE, Sharp C, Petrella RJ. Scoping review report: obesity in older adults. International Journal of Obesity (2005). 2012; 36: 1141–1150.
- [4] Gill LE, Bartels SJ, Batsis JA. Weight Management in Older Adults. Current Obesity Reports. 2015; 4: 379–388.
- [5] Hoseini Tavassol Z, Ejtahed HS, Atlasi R, Saghafian F, Khalagi K, Hasani-Ranjbar S, et al. Alteration in Gut Microbiota Composition of Older Adults Is Associated with Obesity and Its Indices: A Systematic Review. The Journal of Nutrition, Health & Aging. 2023; 27: 817–823.
- [6] Vallianou N, Stratigou T, Christodoulatos GS, Tsigalou C, Dalamaga M. Probiotics, Prebiotics, Synbiotics, Postbiotics, and Obesity: Current Evidence, Controversies, and Perspectives. Current Obesity Reports. 2020; 9: 179–192.
- [7] Setbo E, Campbell K, O'Cuiv P, Hubbard R. Utility of Probiotics for Maintenance or Improvement of Health Status in Older People A Scoping Review. The Journal of Nutrition, Health & Aging. 2019; 23: 364–372.
- [8] Page MJ, McKenzie JE, Bossuyt PM, Boutron I, Hoffmann TC, Mulrow CD, et al. The PRISMA 2020 statement: an updated guideline for reporting systematic reviews. BMJ (Clinical Research Ed.). 2021; 372: n71.
- [9] Higgins JPT, Altman DG, Gøtzsche PC, Jüni P, Moher D, Oxman AD, et al. The Cochrane Collaboration's tool for assessing risk of bias in randomised trials. BMJ (Clinical Research Ed.). 2011; 343: d5928.
- [10] Neto JV, de Melo CM, Ribeiro SML. Effects of three-month intake of synbiotic on inflammation and body composition in the elderly: a pilot study. Nutrients. 2013; 5: 1276–1286.
- [11] Barreto FM, Colado Simão AN, Morimoto HK, Batisti Lozovoy MA, Dichi I, Helena da Silva Miglioranza L. Beneficial effects

- of Lactobacillus plantarum on glycemia and homocysteine levels in postmenopausal women with metabolic syndrome. Nutrition (Burbank, Los Angeles County, Calif.). 2014; 30: 939–942.
- [12] Ivey KL, Hodgson JM, Kerr DA, Lewis JR, Thompson PL, Prince RL. The effects of probiotic bacteria on glycaemic control in overweight men and women: a randomised controlled trial. European Journal of Clinical Nutrition. 2014; 68: 447–452.
- [13] Brahe LK, Le Chatelier E, Prifti E, Pons N, Kennedy S, Blædel T, *et al.* Dietary modulation of the gut microbiota—a randomised controlled trial in obese postmenopausal women. The British Journal of Nutrition. 2015; 114: 406–417.
- [14] Tajabadi-Ebrahimi M, Sharifi N, Farrokhian A, Raygan F, Karamali F, Razzaghi R, et al. A Randomized Controlled Clinical Trial Investigating the Effect of Synbiotic Administration on Markers of Insulin Metabolism and Lipid Profiles in Overweight Type 2 Diabetic Patients with Coronary Heart Disease. Experimental and Clinical Endocrinology & Diabetes: Official Journal, German Society of Endocrinology [and] German Diabetes Association. 2017; 125: 21–27.
- [15] Raygan F, Ostadmohammadi V, Bahmani F, Asemi Z. The effects of vitamin D and probiotic co-supplementation on mental health parameters and metabolic status in type 2 diabetic patients with coronary heart disease: A randomized, double-blind, placebo-controlled trial. Progress in Neuropsychopharmacology & Biological Psychiatry. 2018; 84: 50– 55.
- [16] Szulińska M, Łoniewski I, van Hemert S, Sobieska M, Bog-dański P. Dose-Dependent Effects of Multispecies Probiotic Supplementation on the Lipopolysaccharide (LPS) Level and Cardiometabolic Profile in Obese Postmenopausal Women: A 12-Week Randomized Clinical Trial. Nutrients. 2018; 10: 773.
- [17] Cancello R, Turroni S, Rampelli S, Cattaldo S, Candela M, Cattani L, et al. Effect of Short-Term Dietary Intervention and Probiotic Mix Supplementation on the Gut Microbiota of Elderly Obese Women. Nutrients. 2019; 11: 3011.
- [18] Farrokhian A, Raygan F, Soltani A, Tajabadi-Ebrahimi M, Sharifi Esfahani M, Karami AA, et al. The Effects of Synbiotic Supplementation on Carotid Intima-Media Thickness, Biomarkers of Inflammation, and Oxidative Stress in People with Overweight, Diabetes, and Coronary Heart Disease: a Randomized, Double-Blind, Placebo-Controlled Trial. Probiotics and Antimicrobial Proteins. 2019; 11: 133–142.
- [19] Raji Lahiji M, Najafi S, Janani L, Yazdani B, Razmpoosh E, Zarrati M. The effect of synbiotic on glycemic profile and sex hormones in overweight and obese breast cancer survivors following a weight-loss diet: A randomized, triple-blind, controlled trial. Clinical Nutrition (Edinburgh, Scotland). 2021; 40: 394–403.
- [20] Chaiyasut C, Sivamaruthi BS, Lailerd N, Sirilun S, Khongtan S, Fukngoen P, et al. Probiotics Supplementation Improves Intestinal Permeability, Obesity Index and Metabolic Biomarkers in Elderly Thai Subjects: A Randomized Controlled Trial. Foods (Basel, Switzerland). 2022; 11: 268.
- [21] Kaczmarczyk M, Szulińska M, Łoniewski I, Kręgielska-Narożna M, Skonieczna-Żydecka K, Kosciolek T, et al. Treatment With Multi-Species Probiotics Changes the Functions, Not the Composition of Gut Microbiota in Postmenopausal Women With Obesity: A Randomized, Double-Blind, Placebo-Controlled Study. Frontiers in Cellular and Infection Microbiology. 2022; 12: 815798.
- [22] Brummer R. Effect of Omega-3 and Probiotic Dietary Supplements on Elevated High Sensitivity C-reactive Protein (Hs-CRP) as a Marker of Low-grade Inflammation: Targeting Inflammation and Intestinal Barrier Function in the Elderly and Obese. In: Örebro University S, Hohenheim Uo, Pfizer, editors.: clinicaltrials.gov; 2021.

- [23] Borgeraas H, Johnson LK, Skattebu J, Hertel JK, Hjelmesaeth J. Effects of probiotics on body weight, body mass index, fat mass and fat percentage in subjects with overweight or obesity: a systematic review and meta-analysis of randomized controlled trials. Obesity Reviews: an Official Journal of the International Association for the Study of Obesity. 2018; 19: 219–232.
- [24] Wang ZB, Xin SS, Ding LN, Ding WY, Hou YL, Liu CQ, et al. The Potential Role of Probiotics in Controlling Overweight/Obesity and Associated Metabolic Parameters in Adults: A Systematic Review and Meta-Analysis. Evidence-based Complementary and Alternative Medicine: ECAM. 2019; 2019: 3862971.
- [25] Álvarez-Arraño V, Martín-Peláez S. Effects of Probiotics and Synbiotics on Weight Loss in Subjects with Overweight or Obesity: A Systematic Review. Nutrients. 2021; 13: 3627.
- [26] Shirvani-Rad S, Tabatabaei-Malazy O, Mohseni S, Hasani-Ranjbar S, Soroush AR, Hoseini-Tavassol Z, et al. Probiotics as a Complementary Therapy for Management of Obesity: A Systematic Review. Evidence-based Complementary and Alternative Medicine: ECAM. 2021; 2021: 6688450.
- [27] Tomé-Castro XM, Rodriguez-Arrastia M, Cardona D, Rueda-Ruzafa L, Molina-Torres G, Roman P. Probiotics as a therapeutic strategy in obesity and overweight: a systematic review. Beneficial Microbes. 2021; 12: 5–15.
- [28] Tam BT, Morais JA, Santosa S. Obesity and ageing: Two sides of the same coin. Obesity Reviews: an Official Journal of the International Association for the Study of Obesity. 2020; 21: e12991.
- [29] Bernhold QS, Giles H. Older Adults' Age-Related Communication and Routine Dietary Habits. Health Communication. 2020; 35: 1556–1564.
- [30] Silveira EA, Mendonça CR, Delpino FM, Elias Souza GV, Pereira de Souza Rosa L, de Oliveira C, et al. Sedentary behavior, physical inactivity, abdominal obesity and obesity in adults and older adults: A systematic review and meta-analysis. Clinical Nutrition ESPEN. 2022; 50: 63–73.
- [31] López-Moreno A, Suárez A, Avanzi C, Monteoliva-Sánchez M, Aguilera M. Probiotic Strains and Intervention Total Doses for Modulating Obesity-Related Microbiota Dysbiosis: A Systematic Review and Meta-analysis. Nutrients. 2020; 12: 1921.
- [32] Crovesy L, Ostrowski M, Ferreira DMTP, Rosado EL, Soares-Mota M. Effect of Lactobacillus on body weight and body fat in overweight subjects: a systematic review of randomized controlled clinical trials. International Journal of Obesity (2005). 2017; 41: 1607–1614.
- [33] Ejtahed HS, Angoorani P, Soroush AR, Atlasi R, Hasani-Ranjbar S, Mortazavian AM, et al. Probiotics supplementation for the obesity management; A systematic review of animal studies and clinical trials. Journal of Functional Foods. 2019; 52: 228–242.
- [34] Kim J, Yun JM, Kim MK, Kwon O, Cho B. Lactobacillus gasseri BNR17 Supplementation Reduces the Visceral Fat Accumulation and Waist Circumference in Obese Adults: A Randomized, Double-Blind, Placebo-Controlled Trial. Journal of Medicinal Food. 2018; 21: 454–461.
- [35] Sanchez M, Darimont C, Drapeau V, Emady-Azar S, Lepage M, Rezzonico E, et al. Effect of Lactobacillus rhamnosus CGMCC1.3724 supplementation on weight loss and maintenance in obese men and women. The British Journal of Nutrition. 2014; 111: 1507–1519.
- [36] Liu YW, Liong MT, Tsai YC. New perspectives of Lactobacillus plantarum as a probiotic: The gut-heart-brain axis. Journal of Microbiology (Seoul, Korea). 2018; 56: 601–613.
- [37] Mo SJ, Lee K, Hong HJ, Hong DK, Jung SH, Park SD, et al. Effects of Lactobacillus curvatus HY7601 and Lactobacillus plantarum KY1032 on Overweight and the Gut Microbiota in Humans: Randomized, Double-Blinded, Placebo-Controlled Clin

- ical Trial. Nutrients. 2022; 14: 2484.
- [38] Drissi F, Merhej V, Angelakis E, El Kaoutari A, Carrière F, Henrissat B, *et al.* Comparative genomics analysis of Lactobacillus species associated with weight gain or weight protection. Nutrition & Diabetes. 2014; 4: e109.
- [39] Mao K, Gao J, Wang X, Li X, Geng S, Zhang T, et al. Bifi-dobacterium animalis subsp. lactis BB-12 Has Effect Against Obesity by Regulating Gut Microbiota in Two Phases in Human Microbiota-Associated Rats. Frontiers in Nutrition. 2022; 8: 811619.
- [40] Stenman LK, Lehtinen MJ, Meland N, Christensen JE, Yeung N, Saarinen MT, et al. Probiotic With or Without Fiber Controls Body Fat Mass, Associated With Serum Zonulin, in Overweight and Obese Adults-Randomized Controlled Trial. EBioMedicine. 2016; 13: 190–200.
- [41] Pedret A, Valls RM, Calderón-Pérez L, Llauradó E, Companys J, Pla-Pagà L, *et al.* Effects of daily consumption of the probiotic Bifidobacterium animalis subsp. lactis CECT 8145 on anthropometric adiposity biomarkers in abdominally obese subjects: a randomized controlled trial. International Journal of Obesity (2005). 2019; 43: 1863–1868.
- [42] Uusitupa HM, Rasinkangas P, Lehtinen MJ, Mäkelä SM, Airaksinen K, Anglenius H, et al. Bifidobacterium animalis subsp. lactis 420 for Metabolic Health: Review of the Research. Nutrients. 2020; 12: 892.
- [43] Rahman MS, Lee Y, Park DS, Kim YS. Bifidobacterium bi-fidum DS0908 and Bifidobacterium longum DS0950 Culture-Supernatants Ameliorate Obesity-Related Characteristics in Mice with High-Fat Diet-Induced Obesity. Journal of Microbiology and Biotechnology. 2023; 33: 96–105.
- [44] Solito A, Bozzi Cionci N, Calgaro M, Caputo M, Vannini L, Hasballa I, *et al.* Supplementation with Bifidobacterium breve BR03 and B632 strains improved insulin sensitivity in chil-

- dren and adolescents with obesity in a cross-over, randomized double-blind placebo-controlled trial. Clinical Nutrition (Edinburgh, Scotland). 2021; 40: 4585–4594.
- [45] Wiciński M, Gębalski J, Gołębiewski J, Malinowski B. Probiotics for the Treatment of Overweight and Obesity in Humans-A Review of Clinical Trials. Microorganisms. 2020; 8: 1148.
- [46] Crovesy L, Masterson D, Rosado EL. Profile of the gut microbiota of adults with obesity: a systematic review. European Journal of Clinical Nutrition. 2020; 74: 1251–1262.
- [47] Michael DR, Jack AA, Masetti G, Davies TS, Loxley KE, Kerry-Smith J, et al. A randomised controlled study shows supplementation of overweight and obese adults with lactobacilli and biffdobacteria reduces bodyweight and improves well-being. Scientific Reports. 2020; 10: 4183.
- [48] Michael DR, Davies TS, Jack AA, Masetti G, Marchesi JR, Wang D, et al. Daily supplementation with the Lab4P probiotic consortium induces significant weight loss in overweight adults. Scientific Reports. 2021; 11: 5.
- [49] Rytter H, Combet E, Chassaing B. Probiotic: is diet part of the efficacy equation? Gut Microbes. 2023; 15: 2222438.
- [50] Purdel C, Ungurianu A, Adam-Dima I, Margină D. Exploring the potential impact of probiotic use on drug metabolism and efficacy. Biomedicine & Pharmacotherapy. 2023; 161: 114468.
- [51] Vemuri RC, Gundamaraju R, Shinde T, Eri R. Therapeutic interventions for gut dysbiosis and related disorders in the elderly: antibiotics, probiotics or faecal microbiota transplantation? Beneficial Microbes. 2017; 8: 179–192.
- [52] Hutchinson AN, Bergh C, Kruger K, Sűsserová M, Allen J, Améen S, et al. The Effect of Probiotics on Health Outcomes in the Elderly: A Systematic Review of Randomized, Placebo-Controlled Studies. Microorganisms. 2021; 9: 1344.
- [53] Chaudhary P, Kathuria D, Suri S, Bahndral A, Naveen AK. Probiotics-its functions and influence on the ageing process: A comprehensive review. Food Bioscience. 2023; 52: 102389.